

Biostatistics 202: Logistic regression analysis

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In our last article on linear regression⁽¹⁾, we modeled the relationship between the systolic blood pressure, which was a continuous quantitative outcome, with age, race and smoking status of 55 subjects. If our interest now is to model the predictors for SBP ≥ 180 mmHg, a categorical dichotomous outcome (Table I), then the appropriate multivariate analysis is a logistic regression.

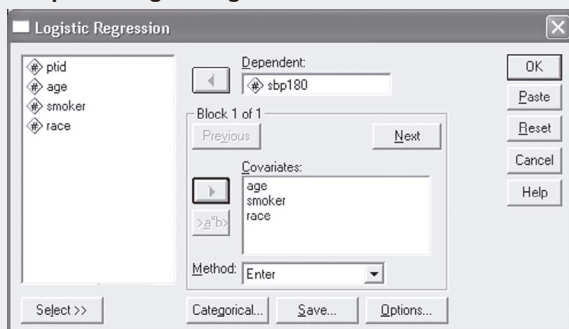
Table I. Frequency distribution of SBP ≥ 180 mmHg.

| | | sbp ≥ 180 | | | |
|-------|-----|----------------|---------|---------------|--------------------|
| | | Frequency | Percent | Valid percent | Cumulative percent |
| Valid | no | 40 | 72.7 | 72.7 | 72.7 |
| | yes | 15 | 27.3 | 27.3 | 100.00 |
| Total | | 55 | 100.0 | 100.0 | |

Since our interest is to determine the predictors for SBP ≥ 180 mmHg, then the numerical coding for SBP ≥ 180 mmHg must be “bigger” than that of SBP < 180 mmHg, say 1 & 0, respectively. SPSS will use the “higher coded” category to be the predicted outcome.

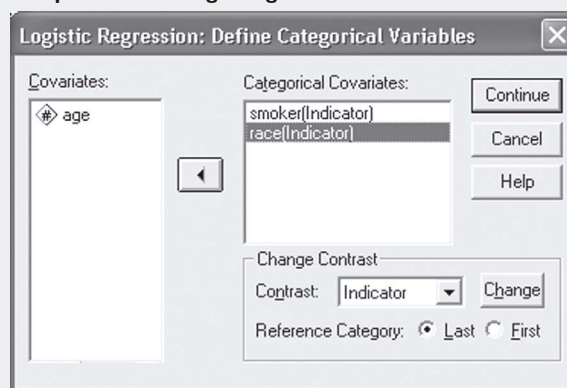
To perform the logistic regression using SPSS, go to *Analyze, Regression, Binary Logistic* to get template I.

Template I. Logistic regression.



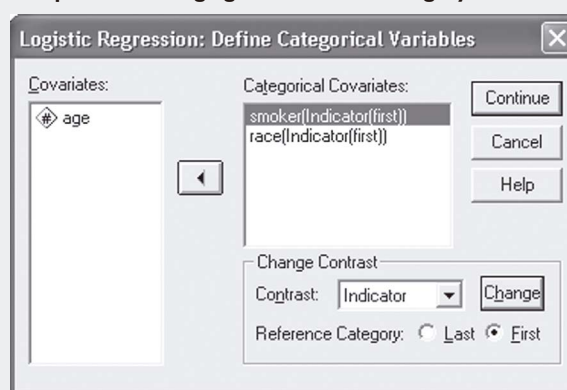
Put sbp180 (the categorized SBP ≥ 180 mmHg & SBP < 180 mmHg) in the Dependent box. Put age, race and smoker in the Covariates box. Click on the Categorical folder (in template I) to declare smoker and race as categorical variables (Template II).

Template II. Defining categorical variables.



Since smoker and race are categorical, we will need a reference group (the default is the “highest coded” Last category). For race, usually we want the Chinese to be the reference and our standard coding is 1 = Chinese, 2 = Indian, 3 = Malay, 4 = Others, then we got to change the Reference Category (at the bottom of template II) to First and click on the Change button (Template III).

Template III. Changing the reference category.



Likewise, we have also changed the reference category for smoking to First as the coding is 1 = smoker and 0 = non smoker. The idea is to prepare the output for “easy interpretation”; that is, comparing the smoker with the non-smoker of having SBP ≥ 180 . Tables IIa – IIe (only those of interest) are the output generated by SPSS when a logistic regression is performed.

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Table IIa. Number of cases in model.

Case processing summary

| Unweighted Cases ^a | | N | Percent |
|-------------------------------|----------------------|----|---------|
| Selected cases | Included in analysis | 55 | 100.0 |
| | Missing cases | 0 | .0 |
| | Total | 55 | 100.0 |
| Unselected cases | | 0 | .0 |
| Total | | 55 | 100.0 |

^a If weight is in effect, see classification table for the total number of cases.

All 55 cases were included in the analysis. A subject will be omitted from the analysis if any one of his data point (for example, age) is missing, regardless of the availability of the others.

Table IIb. Predicted outcome coding.

Dependent variable encoding

| Original value | Internal value |
|----------------|----------------|
| No | 0 |
| Yes | 1 |

Table IIb is very important. It tells us which category SPSS is using as the predicted outcome, the higher coded category (having SBP ≥180 mmHg).

Table IIc. Amount of variation explained by the model.

Model summary

| Step | -2 Log likelihood | Cox & Snell R Square | Nagelkerke R Square |
|------|-------------------|----------------------|---------------------|
| 1 | 40.819 | .349 | .506 |

The Nagelkerke R Square shows that about 50% of the variation in the outcome variable (SBP ≥180) is explained by this logistic model.

Table IIId. Estimates of the logistic regression model.

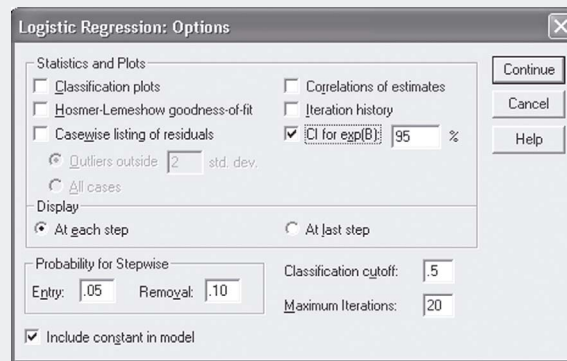
Variables in the equation

| | | B | S.E. | Wald | df | Sig. | Exp(B) | 95.0% C.I. for EXP(B) | |
|---------------------|-----------|---------|-------|--------|----|------|--------|-----------------------|--------|
| | | | | | | | | Lower | Upper |
| Step 1 ^a | AGE | .209 | .063 | 11.007 | 1 | .001 | 1.233 | 1.089 | 1.395 |
| | SMOKER(1) | 2.292 | .986 | 5.401 | 1 | .020 | 9.896 | 1.432 | 68.380 |
| | RACE | | | 1.627 | 3 | .653 | | | |
| | RACE(1) | .640 | 1.009 | .402 | 1 | .526 | 1.896 | .263 | 13.696 |
| | RACE(2) | 1.303 | 1.136 | 1.316 | 1 | .251 | 3.681 | .397 | 34.101 |
| | RACE(3) | -.097 | 1.230 | .006 | 1 | .937 | .908 | .081 | 10.113 |
| | Constant | -14.462 | 4.005 | 13.041 | 1 | .000 | .000 | | |

How do we interpret the results in Table IIId? Firstly, the **Wald** estimates give the “importance” of the contribution of each variable in the model. The higher the value, the more “important” it is.

If we are interested in a **predictor-model**, then both age and smoking status are important risk factors to having SBP ≥180, with p-values of 0.001 and 0.020 (given by the Sig column), respectively. The **Exp(B)** gives the **Odds Ratios**. Since age is a quantitative numerical variable, an increase in one-year in age has a 23.3% (95% CI 8.9% to 39.5%) increase in odds of having SBP ≥180. This 23.3% is obtained by taking Exp(B) for age – 1. To get the 95% CI, in Template I, click on the Options folder to get Template IV.

Template IV. Getting the 95% CI for the odds ratios.



Tick on CI for exp(B) for the 95% CI of the estimate.

In Table IIId, what is SMOKER(1)? Table IIe shows the coding for the categorical variables. The reference group for a particular variable is given by the row of zeros. Thus for Smoker, the reference group is the non-smoker (as setup in Template III). A smoker compared to a non-smoker is 9.9 (95% CI 1.4 to 68.4) times more likely to have SBP ≥180.

Table IIe. Categorical variables coding.

| Categorical variables codings | | | Parameter coding | | |
|-------------------------------|---------|----|------------------|-------|-------|
| | | | Frequency | (1) | (2) |
| RACE | Chinese | 23 | .000 | .000 | .000 |
| | Indian | 13 | 1.000 | .000 | .000 |
| | Malay | 10 | .000 | 1.000 | .000 |
| | Others | 9 | .000 | .000 | 1.000 |
| Smoker | No | 23 | .000 | | |
| | Yes | 32 | 1.000 | | |

For Race, Chinese is the reference category. In Table IIId, Race(1) refers to comparing the Indian with Chinese, Race(2) refers to comparing the Malay with Chinese and lastly, Race(3) for Others comparing with Chinese. In Template III, observe that we can only declare either the first or last as the reference. If we want Malay to be the reference, a recode to make Malay having the smallest or largest coding is required.

CHECKING MULTICOLINEARITY

How to check for multicollinearity? To get the correlations between any two variables, in Template IV, tick on the Correlations of estimates option to obtain table III.

Table III. Correlation matrix for SBP model.

| | | Correlation matrix | | | | | |
|--------|-----------|--------------------|-----------|---------|---------|---------|-------|
| | | Constant | SMOKER(1) | RACE(1) | RACE(2) | RACE(3) | AGE |
| Step I | Constant | 1.000 | .345 | -.326 | -.265 | -.415 | -.953 |
| | SMOKER(1) | .345 | 1.000 | .073 | .081 | -.122 | -.450 |
| | RACE(1) | -.326 | .073 | 1.000 | .700 | .652 | .068 |
| | RACE(2) | -.265 | .081 | .700 | 1.000 | .585 | .030 |
| | RACE(3) | -.415 | -.122 | .652 | .585 | 1.000 | .215 |
| | AGE | -.953 | -.450 | .068 | .030 | .215 | 1.000 |

Apart from the expected moderate to high correlations within Race, the correlation values among age, smoker and race are low. The correlation between age and the constant is rather high ($r = -0.953$) which shows some multicollinearity. What should be done? Before we answer this question, let us look at another example which quite commonly happens in a many-variables study. Table IV shows a 8-variable model with the correlation matrix between any two variables given in Table V.

Table IV. An 8-variable logistic model with multicollinearity.

| | | Variables in the Equation | | | | |
|----------|----|---------------------------|------------|------|----|------|
| | | B | S.E. | Wald | df | Sig. |
| Step I | V1 | -1062.640 | 56906.272 | .000 | 1 | .985 |
| | V2 | -2033.243 | 107665.309 | .000 | 1 | .985 |
| | V3 | -2282.536 | 121116.943 | .000 | 1 | .985 |
| | V4 | -462.334 | 26296.043 | .000 | 1 | .986 |
| | V5 | 1000.935 | 53615.449 | .000 | 1 | .985 |
| | V6 | 65.543 | 5358.046 | .000 | 1 | .990 |
| | V7 | 764.889 | 40207.609 | .000 | 1 | .985 |
| | V8 | -62.261 | 4286.793 | .000 | 1 | .988 |
| Constant | | -829.405 | 44003.539 | .000 | 1 | .985 |

In the correlation matrix for this case, it is not so easy to spot where the multicollinearity is! Another drawback with the correlation matrix is that multicollinearity between one variable with a combination of variables will not be shown.

A simple but sometimes subjective technique is to inspect the magnitude of the standard error (SE) of each variable. The SEs in Table IV are very large implying multicollinearity exists and the model is not statistically stable. To “solve” this issue, start omitting the variable with largest SE, continue the process until the magnitude of the SEs hover around 0.001 – 5.0. There is no fixed criterion on how small the SE should be but a matter of judgment.

In Table IIId, the SEs are within the acceptable criterion but there was a high correlation between age and the constant – should one of them be omitted? The recommendation is to keep the constant term in the model as it acts as a “garbage bin”, collecting all unexplained variance in the model (recall from Table IIc that the variables only explains 50%). How to omit the constant? In template IV, at the left hand corner, uncheck the “Include constant in model”.

A PREDICTION MODEL

Frequently our interest is to use the logistic model to predict the outcome for a new subject. How good is this model for prediction?

Table V. Correlation matrix of the 8-variable model.

| | | Correlation matrix | | | | | | | | |
|--------|----------|--------------------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | Constant | V1 | V2 | V3 | V4 | V5 | V6 | V7 | V8 |
| Step I | Constant | 1.000 | -.878 | -.892 | .965 | -.920 | -.924 | -.917 | -.523 | -.412 |
| | V1 | -.878 | 1.000 | .659 | -.831 | .743 | .938 | .766 | .144 | .389 |
| | V2 | -.892 | .659 | 1.000 | -.887 | .866 | .746 | .809 | .679 | .222 |
| | V3 | .965 | -.831 | -.887 | 1.000 | -.980 | -.917 | -.887 | -.555 | -.374 |
| | V4 | -.920 | .743 | .866 | -.980 | 1.000 | .877 | .832 | .598 | .342 |
| | V5 | -.924 | .938 | .746 | -.917 | .877 | 1.000 | .799 | .280 | .378 |
| | V6 | -.917 | .766 | .809 | -.887 | .832 | .799 | 1.000 | .620 | .150 |
| | V7 | -.523 | .144 | .679 | -.555 | .598 | .280 | .620 | 1.000 | -.155 |
| | V8 | -.412 | .389 | .222 | -.374 | .342 | .378 | .150 | -.155 | 1.000 |

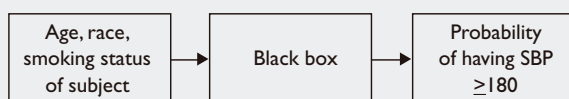
Table VI. Model discrimination.

| Classification table ^a | | | | | |
|-----------------------------------|----------------|----------------|-----|--------------------|------|
| | | Predicted | | | |
| | | SBP \geq 180 | | Percentage correct | |
| Observed | | no | yes | | |
| Step I | SBP \geq 180 | no | 38 | 2 | 95.0 |
| | | yes | 6 | 9 | 60.0 |
| Overall percentage | | | | | 85.5 |

^a The cut value is .500

The overall accuracy of this model to predict subjects having SBP \geq 180 (with a predicted probability of 0.5 or greater) is 85.5% (Table VI). The sensitivity is given by $9/15 = 60\%$ and the specificity is $38/40 = 95\%$. Positive predictive value (PPV) = $9/11 = 81.8\%$ and negative predictive value (NPV) = $38/44 = 86.4\%$. How to use this information?

When we have a new subject, we can use the logistic model to predict his probability of having SBP \geq 180. Let us say we have a black box where we input the age, smoking status and race of a subject and the output is a number between 0 to 1 which denotes the probability of the subject having SBP \geq 180 (see Fig. 1).

Fig. 1 The logistic regression prediction model.

In the black box, we have the equation for calculating the probability of having SBP \geq 180 which is given by

$$\text{Prob (SBP} \geq 180) = \frac{1}{1+e^{-z}}$$

where e denotes the exponential function

$$\text{with } z = -14.462 + 0.209 * \text{Age} + 2.292 * \text{Smoker}(1) + 0.640 * \text{Race}(1) + 1.303 * \text{Race}(2) - 0.097 * \text{Race}(3)$$

The numerical values are obtained from the B estimates in Table II.

For example, we have a 45-year-old non-smoking Chinese, then Smoker(1) = Race(2) = Race(3) = 0, and

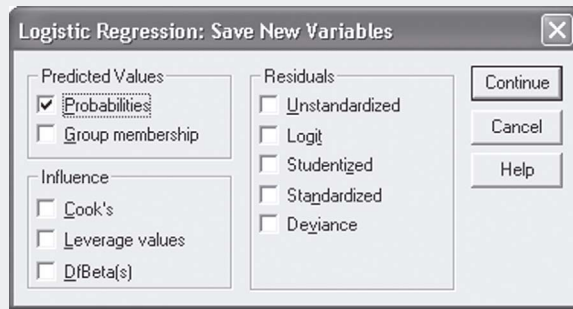
$z = -14.462 + 0.209 * 45 = -5.057$ and $e^{-z} = 157.1$ which gives the Prob (SBP \geq 180) = $1 / (1 + 157.1) = 0.006$; very unlikely that this subject has SBP \geq 180 and the NPV tells me that I am 86.4% confident.

Let us take another example, a 65-year-old Indian smoker, then Smoker(1) = 1, Race(2) = Race(3) = 0 but Race(1) = 1. Hence $z = -14.462 + 0.209 * 65 + 2.292 * 1 + 0.64 * 1 = 2.055$ and $e^{-z} = 0.128$ which gives the Prob (SBP \geq 180) = $1 / (1 + 0.128) = 0.89$; very likely that this subject has SBP \geq 180 and the PPV gives a 81.8% confidence.

The default cut-off probability is 0.5 (and for this model, it seems that this cut-off gives quite good results). We can generate different probability cutoffs, by changing the 'Classification cutoff' in Template IV, and tabulate the respective sensitivity, specificity, PPV and NPV, then decide which is the best cut-off for optimal results.

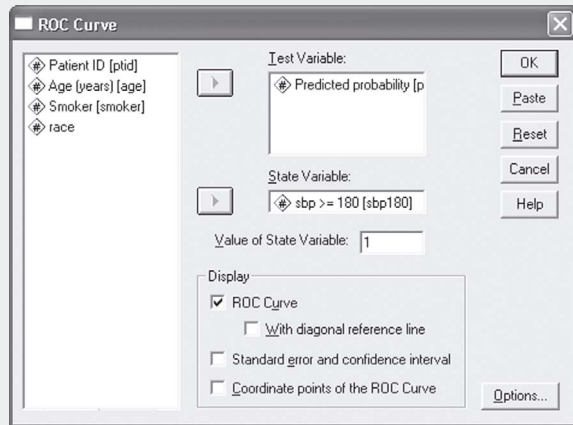
The **area under the ROC curve**, which ranges from 0 to 1, could also be used to assess the model discrimination. A value of 0.5 means that the model is useless for discrimination (equivalent to tossing a coin) and values near 1 means that higher probabilities will be assigned to cases with the outcome of interest compared to cases without the outcome. To generate the ROC, we have to save the predicted probabilities from the model. In Template I, click on the Save button to get Template V.

Template V. Saving the predicted probabilities.



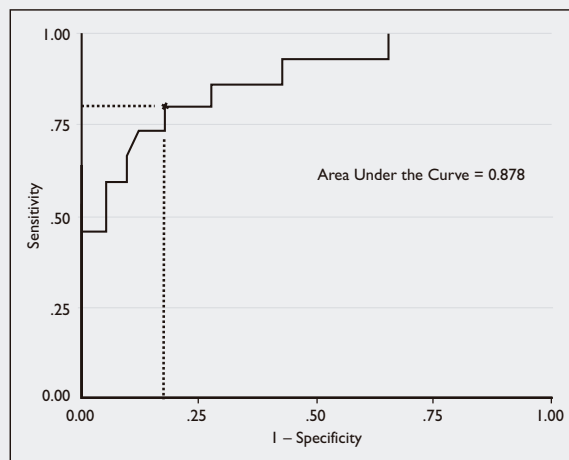
Check the Predicted Values – Probabilities. A new variable, pre_1 (Predicted probability), will be created when the logistic regression is performed. Next go to **Graphs, ROC curve** – see Template VI.

Template VI. ROC curve.



Put Predicted probability (pre_1) into the test Variable box, sbp180 in the State Variable and Value of State Variable = 1 (to predict SBP ≥180).

Fig. 2 ROC curve and area.



The ROC area is 0.878 (Fig. 2) which means that in almost 88% of all possible pairs of subjects in which one has SBP ≥180 and the other SBP <180, this model will assign a higher probability to the subject with SBP ≥180. The optimal sensitivity/specificity is obtained from the point (*) nearest to the left upper corner of the box. Thus the optimal sensitivity = 78% and specificity = 1 - 0.18 = 82%.

Hosmer-Lemeshow goodness of fit (obtained by checking the relevant box in template IV) tells us how closely the observed and predicted probabilities match. The null hypothesis is “the model fits” and a p value >0.05 is expected (Table VII). Caution has to be exercised when using this test as it is dependent on the sample size of the data. For a small sample size, this test will likely indicate that the model fits and for a large dataset, even if the model fits, this test may “fail”.

Table VII. Hosmer-Lemeshow test.

| Hosmer and Lemeshow Test | | | |
|--------------------------|------------|----|------|
| Step | Chi-square | df | Sig. |
| 1 | 5.869 | 7 | .555 |

The above material covered the situation where the response outcome has only two levels. There are times when it is not possible to collapse the outcome of interest into two groups, for example stage of cancer. There are also situations where our study is a matched case-control. If in doubt, do seek help from a Biostatistician. The next article, Biostatistics 203, will be on Survival Analysis.

REFERENCE

1. Chan YH, Biostatistics 201: Linear regression analysis. Singapore Med J 2004; 45:55-61.